

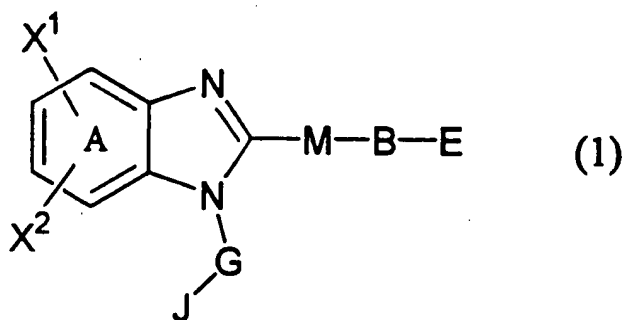
**AMENDMENTS TO THE CLAIMS**

**This listing of claims will replace all prior versions and listings of claims in the application:**

**LISTING OF CLAIMS:**

**1-21. (Canceled)**

**22. (New):** A method of inhibiting human chymase activity to treat or prevent a bone/cartilage metabolic disease in human beings, said method comprising administering to a subject an effective amount of a benzimidazole derivative expressed by the following formula



(1) or its pharmaceutically permissible salt,

[in the formula (1), the ring marked with A expresses a benzene ring;

X<sup>1</sup> and X<sup>2</sup> are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a hydroxyl group, a nitro group, a cyano group, -CH<sub>2</sub>NH<sub>2</sub>, -CH=NR<sup>1</sup>, -CH=NOR<sup>1</sup> or -CONR<sup>1</sup>R<sup>2</sup> (here, R<sup>1</sup> and R<sup>2</sup> are each a hydrogen atom or a C<sub>1-4</sub> alkyl group), -COOR<sup>3</sup> (here, R<sup>3</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkyl group, a substituted or unsubstituted C<sub>3-7</sub> cycloalkyl group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkoxy group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylthio group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched

alkylsulfonyl group or a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylsulfinyl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s)};

B is a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkylene group or a substituted or unsubstituted C<sub>2-6</sub> normal or branched alkenylene group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a C<sub>1-6</sub> normal or branched alkoxyl group (including the case where adjacent two groups form an acetal bonding), a C<sub>1-6</sub> normal or branched alkylthio group, a C<sub>1-6</sub> normal or branched alkylsulfonyl group, a C<sub>1-6</sub> normal or branched acyl group, a C<sub>1-6</sub> normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s) of the alkylene group or an alkenylene group; between atoms, the alkylene group or alkenylene group optionally contains one or more of -O-, -S-, -SO<sub>2</sub>- or -NR<sup>4</sup>-, but this atom or atomic group does not bond directly to the M, and here R<sup>4</sup> is a hydrogen atom or a C<sub>1-6</sub> normal or branched alkyl group};

E expresses -COOR<sup>4</sup>;

G is a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylene group {between atoms, the alkylene group optionally contains one or more of -O-, -S-, -SO<sub>2</sub>- or -NR<sup>4</sup>-, but this atom or atomic group does not bond directly to the nitrogen atom of the imidazole ring (R<sup>4</sup> is similarly defined as above), and the substituent is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a C<sub>1-6</sub> normal or branched alkoxyl group (including the case where

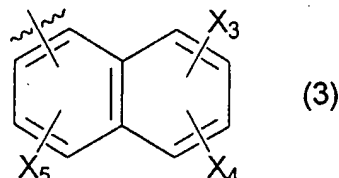
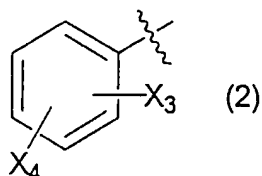
adjacent two groups form an acetal bonding), a trihalomethyl group, a trihalomethoxy group, a phenyl group or an oxo group};

J is a substituted or unsubstituted C<sub>4-10</sub> aryl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, -COOR<sup>7</sup> (here, R<sup>7</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a C<sub>1-6</sub> normal, cyclic or branched alkyl group, a C<sub>1-6</sub> normal or branched alkoxy group (including the case where adjacent two groups form an acetal bonding), a C<sub>1-6</sub> normal or branched alkylthio group, a C<sub>1-6</sub> normal or branched alkylsulfonyl group, a C<sub>1-6</sub> normal or branched alkylsulfinyl group, a C<sub>1-6</sub> acyl group, a C<sub>1-6</sub> normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group, or a phenoxy group optionally substituted with one or more halogen atoms; the substituent may substitute singly or plurally independently at arbitrary position(s) of the aryl group; and the substituent is further optionally substituted with a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a phenyl group, an oxo group or a phenoxy group optionally substituted with a halogen atom}; and

M is a sulfur atom, a sulfinyl group, or a sulfonyl group].

**23. (New):** The method set forth in Claim 22 wherein X<sup>1</sup> and X<sup>2</sup> in the above formula (1) are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a cyano group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkyl group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkoxy group, or a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylthio group.

**24. (New):** The method set forth in Claim 22 wherein J in formula (1) is a group described in the following formula (2) or (3),



[here,  $X^3$ ,  $X^4$  and  $X^5$  are each at the same time or independently a hydrogen atom, a halogen atom, a hydroxyl group, a nitro group, a cyano group, a trihalomethyl group, a trihalomethoxy group,  $-\text{COOR}^7$  (here,  $R^7$  is a hydrogen atom or a  $\text{C}_{1-4}$  alkyl group), a substituted or unsubstituted  $\text{C}_{1-3}$  normal or branched alkyl group, a substituted or unsubstituted  $\text{C}_{1-3}$  normal or branched alkoxyl group, a substituted or unsubstituted  $\text{C}_{1-3}$  normal or branched alkylthio group, a substituted or unsubstituted  $\text{C}_{1-3}$  normal or branched alkylsulfonyl group, or a substituted or unsubstituted  $\text{C}_{1-3}$  normal or branched alkylsulfinyl group; there is no limitation regarding the substitution positions of  $X^3$ ,  $X^4$  and  $X^5$  on the benzene ring or the naphthalene ring].

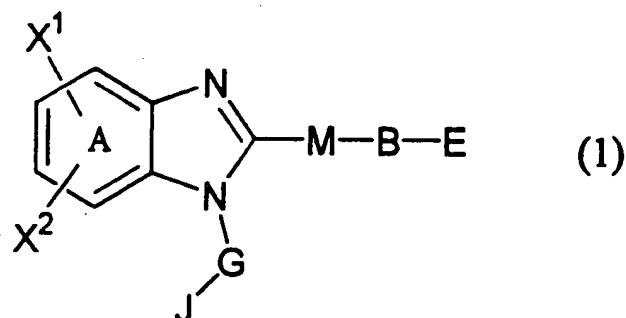
**25. (New):** The method set forth in Claim 22 wherein M is a sulfur atom.

**26. (New):** The method set forth in Claim 22 wherein B is a substituted or unsubstituted  $\text{C}_{1-6}$  normal, cyclic or branched alkylene group.

**27. (New):** The method set forth in Claim 22 wherein G is  $-\text{CH}_2-$ ,  $-\text{CH}_2\text{CH}_2-$ ,  $-\text{CH}_2\text{CO}-$ ,  $-\text{CH}_2\text{CH}_2\text{O}-$ ,  $-\text{CH}_2\text{CONH}-$ ,  $-\text{CO}-$ ,  $-\text{SO}_2-$ ,  $-\text{CH}_2\text{SO}_2-$ ,  $-\text{CH}_2\text{S}-$  or  $-\text{CH}_2\text{CH}_2\text{S}-$  (J bonds to the right side of said group).

**28. (New):** The method set forth in Claim 22 wherein E is  $-\text{COOH}$ .

29. (New): A benzimidazole derivative expressed by the following formula (1) or its pharmaceutically permissible salt,



[in the formula (1), the ring marked with A expresses a benzene ring;

X<sup>1</sup> and X<sup>2</sup> are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a hydroxyl group, a nitro group, a cyano group, -CH<sub>2</sub>NH<sub>2</sub>, -CH=NR<sup>1</sup>, -CH=NOR<sup>1</sup> or -CONR<sup>1</sup>R<sup>2</sup> (here, R<sup>1</sup> and R<sup>2</sup> are each a hydrogen atom or a C<sub>1-4</sub> alkyl group), -COOR<sup>3</sup> (here, R<sup>3</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkyl group, a substituted or unsubstituted C<sub>3-7</sub> cycloalkyl group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkoxy group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylthio group, a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylsulfonyl group or a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylsulfinyl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s)};

B is a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkylene group or a substituted or unsubstituted C<sub>2-6</sub> normal or branched alkenylene group {the substituent

permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a C<sub>1-6</sub> normal or branched alkoxyl group (including the case where adjacent two groups form an acetal bonding), a C<sub>1-6</sub> normal or branched alkylthio group, a C<sub>1-6</sub> normal or branched alkylsulfonyl group, a C<sub>1-6</sub> normal or branched acyl group, a C<sub>1-6</sub> normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s) of the alkylene group or an alkenylene group; between atoms, the alkylene group or alkenylene group optionally contains one or more of -O-, -S- or -SO<sub>2</sub>-, but this atom or atomic group does not bond directly to the M, and here R<sup>4</sup> is a hydrogen atom or a C<sub>1-6</sub> normal or branched alkyl group};

E expresses -COOR<sup>4</sup>;

G is a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylene group {between atoms, the alkylene group optionally contains one or more of -O-, -S-, -SO<sub>2</sub>- or -NR<sup>4</sup>-, but this atom or atomic group does not bond directly to the nitrogen atom of the imidazole ring (R<sup>4</sup> is similarly defined as above), and the substituent is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a C<sub>1-6</sub> normal or branched alkoxyl group (including the case where adjacent two groups form an acetal bonding), a trihalomethyl group, a trihalomethoxy group, a phenyl group or an oxo group};

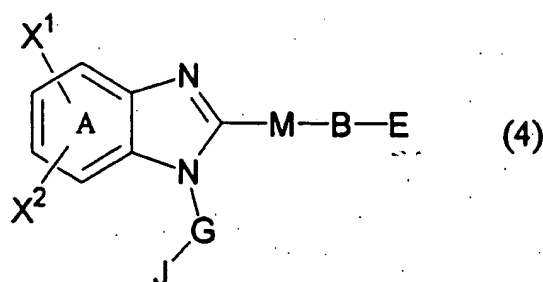
J is a substituted or unsubstituted C<sub>4-10</sub> aryl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, -COOR<sup>7</sup> (here, R<sup>7</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a C<sub>1-6</sub> normal, cyclic or branched alkyl group, a C<sub>1-6</sub> normal or branched alkoxyl group (including the case where adjacent two groups form an acetal bonding), a C<sub>1-6</sub> normal or branched alkylthio group, a C<sub>1-6</sub> normal or branched alkylsulfonyl

group, a C<sub>1-6</sub> normal or branched alkylsulfinyl group, a C<sub>1-6</sub> acyl group, a C<sub>1-6</sub> normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group, or a phenoxy group optionally substituted with one or more halogen atoms; the substituent may substitute singly or plurally independently at arbitrary position(s) of the aryl group; and the substituent is further optionally substituted with a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a phenyl group, an oxo group or a phenoxy group optionally substituted with a halogen atom}; and

M is a sulfinyl group, or a sulfonyl group].

**30. (New):** The benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 wherein X<sup>1</sup> and X<sup>2</sup> in the above formula (1) are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a cyano group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkyl group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkoxyl group, or a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylthio group.

**31. (New):** A benzimidazole derivative expressed by the following formula (4) or its pharmaceutically permissible salt,



[in the formula (4), the ring marked with A expresses a benzene ring;

$X^1$  and  $X^2$  are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a hydroxyl group, a nitro group, a cyano group,  $-CH_2NH_2$ ,  $-CH=NR^1$ ,  $-CH=NOR^1$  or  $-CONR^1R^2$  (here,  $R^1$  and  $R^2$  are each a hydrogen atom or a  $C_{1-4}$  alkyl group),  $-COOR^3$  (here,  $R^3$  is a hydrogen atom or a  $C_{1-4}$  alkyl group), a substituted or unsubstituted  $C_{1-6}$  normal, cyclic or branched alkyl group, a substituted or unsubstituted  $C_{3-7}$  cycloalkyl group, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkoxyl group, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylthio group, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylsulfonyl group or a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylsulfinyl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s)};

B is a substituted or unsubstituted  $C_{1-6}$  normal, cyclic or branched alkylene group or a substituted or unsubstituted  $C_{2-6}$  normal or branched alkenylene group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a  $C_{1-6}$  normal or branched alkoxyl group (including the case where adjacent two groups form an acetal bonding), a  $C_{1-6}$  normal or branched alkylthio group, a  $C_{1-6}$  normal or branched alkylsulfonyl group, a  $C_{1-6}$  normal or branched acyl group, a  $C_{1-6}$  normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s) of the alkylene group or an alkenylene group; between atoms, the alkylene group or alkenylene group optionally contains



one or more of -O-, -S- or -SO<sub>2</sub>-, but this atom or atomic group does not bond directly to the M, and here R<sup>4</sup> is a hydrogen atom or a C<sub>1-6</sub> normal or branched alkyl group};

E expresses -COOR<sup>4</sup>;

G is a substituted or unsubstituted C<sub>1-6</sub> normal or branched alkylene group {between atoms, the alkylene group optionally contains one or more of -O-, -S-, -SO<sub>2</sub>- or -NR<sup>4</sup>-, but this atom or atomic group does not bond directly to the nitrogen atom of the imidazole ring (R<sup>4</sup> is similarly defined as above), and the substituent is a halogen atom, a hydroxyl group, a nitro group, a cyano group, a C<sub>1-6</sub> normal or branched alkoxy group (including the case where adjacent two groups form an acetal bonding), a trihalomethyl group, a trihalomethoxy group, a phenyl group or an oxo group};

J is a substituted or unsubstituted C<sub>4-10</sub> aryl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, -COOR<sup>7</sup> (here, R<sup>7</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a C<sub>1-6</sub> normal, cyclic or branched alkyl group, a C<sub>1-6</sub> normal or branched alkoxy group (including the case where adjacent two groups form an acetal bonding), a C<sub>1-6</sub> normal or branched alkylthio group, a C<sub>1-6</sub> normal or branched alkylsulfonyl group, a C<sub>1-6</sub> normal or branched alkylsulfinyl group, a C<sub>1-6</sub> acyl group, a C<sub>1-6</sub> normal or branched acylamino group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group, or a phenoxy group optionally substituted with one or more halogen atoms; the substituent may substitute singly or plurally independently at arbitrary position(s) of the aryl group; and the substituent is further optionally substituted with a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a phenyl group, an oxo group or a phenoxy group optionally substituted with a halogen atom); and

M is a sulfur atom, a sulfinyl group, or sulfonyl group;

provided that the case is excluded where at least one of  $X^1$  and  $X^2$  is a cyano group,  $-CH_2NH_2$ ,  $-CH=NR^1$ ,  $-CH=NOR^1$  or  $-CONR^1R^2$  (here,  $R^1$  and  $R^2$  are each a hydrogen atom or a  $C_{1-4}$  alkyl group), and J expresses only a substituted naphthalene ring.

**32. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 or 31 wherein  $X^1$  and  $X^2$  are each a hydrogen atom, a cyano group,  $-CH_2NH_2$ ,  $-CH=NR^1$ ,  $-CH=NOR^1$  or  $-CONR^1R^2$  (here,  $R^1$  and  $R^2$  are each a hydrogen atom or a  $C_{1-4}$  alkyl group;  $X^1$  and  $X^2$  are not hydrogen at the same time).

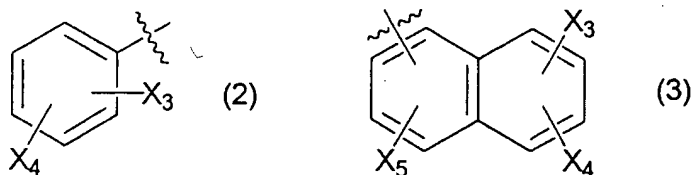
**33. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 or 31 wherein  $X^1$  and  $X^2$  are each at the same time or independently a hydrogen atom, a halogen atom, a trihalomethyl group, a hydroxyl group, a nitro group,  $-COOR^3$  (here,  $R^3$  is a hydrogen atom or a  $C_{1-4}$  alkyl group), a substituted or unsubstituted  $C_{1-6}$  normal, cyclic or branched alkyl group, a substituted or unsubstituted  $C_{3-7}$  cycloalkyl, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkoxyl group, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylthio group, a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylsulfonyl group or a substituted or unsubstituted  $C_{1-6}$  normal or branched alkylsulfinyl group {the substituent permissible to the groups is a halogen atom, a hydroxyl group, a nitro group, a cyano group, an acyl group, a trihalomethyl group, a trihalomethoxy group, a phenyl group, an oxo group or a phenoxy group optionally substituted with one or more halogen atoms, and the substituent may substitute singly or plurally independently at arbitrary position(s)}.

**34. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 or 31 wherein  $X^1$  and  $X^2$  are each a hydrogen atom or a cyano group (here,  $X^1$  and  $X^2$  can not be hydrogen atoms at the same time).

**35. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 31 wherein M is a sulfur atom.

**36. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 or 31 wherein B is a substituted or unsubstituted C<sub>1-6</sub> normal, cyclic or branched alkylene group.

**37. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 or 31 wherein J is a group expressed by the following formula (2) or (3),



[here, X<sup>3</sup>, X<sup>4</sup> and X<sup>5</sup> are each at the same time or independently a hydrogen atom, a halogen atom, a hydroxyl group, a nitro group, a cyano group, a trihalomethyl group, a trihalomethoxy group, -COOR<sup>7</sup> (here, R<sup>7</sup> is a hydrogen atom or a C<sub>1-4</sub> alkyl group), a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkyl group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkoxy group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylthio group, a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylsulfonyl group, or a substituted or unsubstituted C<sub>1-3</sub> normal or branched alkylsulfinyl group; there is no limitation regarding the substitution positions of X<sup>3</sup>, X<sup>4</sup> and X<sup>5</sup> on the benzene ring or the naphthalene ring].

**38. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 or 31 wherein G is -CH<sub>2</sub>-, -CH<sub>2</sub>CH<sub>2</sub>-, -CH<sub>2</sub>CO-, -CH<sub>2</sub>CH<sub>2</sub>O-, -CH<sub>2</sub>CONH-, -CO-, -SO<sub>2</sub>-, -CH<sub>2</sub>SO<sub>2</sub>-, -CH<sub>2</sub>S- or -CH<sub>2</sub>CH<sub>2</sub>S- (J bonds to the right side of said group).

**39. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 29 or 31 wherein E is -COOH.

**40. (New):** A pharmaceutical composition consisting of a benzimidazole derivative and/or its pharmaceutically permissible salt set forth in any one of Claims 29, 30 or 31, and a pharmaceutically permissible carrier.

**41. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in any one of Claims 29, 30 or 31 to prevent or treat an inflammatory disease, an allergy disease, a respiratory disease, a cardiovascular disease or a bone/cartilage metabolic disease.

**42. (New):** A benzimidazole derivative or its pharmaceutically permissible salt set forth in Claim 41 to prevent or treat a disease in human beings.